PATENT COOPERATION TREAT

From the INTERNATIONAL SEARCHING AUTHORITY	PCT	
To: WOLF, GREENFIELD & SACKS, P.C. Attn. Steele, Alan W. 600 Atlantic Avenue	INVITATION TO PAY ADDITIONAL FEES	
Boston, Massachusetts 02210 UNITED STATES OF AMERICA	(PCT Article 17(3)(a) and Rule 40.1)	
	Date of mailing (day/month/year) 29/07/2002	
Applicant's or agent's file reference E0377/7012W0	Within 45 KNSXKS/days	
International application No.	from the above date of mailing International filing date	
PCT/US 01/47251	(day/month/year) 05/12/2001	
Applicant		
THE BRIGHAM AND WOMEN'S HOSPITAL, INC.		
This International Searching Authority		
(i) considers that there are	mber of) inventions claimed in the international application covered Subject to PTA? YES/NO , per docker/ECB)	
and it considers that the international application does not (Rules 13.1, 13.2 and 13.3) for the reasons indicated	Xw/on the extra sheet:	
-	File Follow 105 100 Initials	
	Docket Entry	
A	UG 0 8 2002 Docket Cross Off Order Copies Annuities 0 9 69 65	
I	Confirmation	
(ii) \overline{X} has carried out a partial international search (see Another and those parts of the international application which relate $1-43$		
(iii) will establish the international search report on the other p to which, additional fees are paid	earts of the international application only if, and to the extent	
2. The applicant is hereby invited, within the time limit indicated a	above, to pay the amount indicated below:	
EUR 945, - x 1	= <u>FUR 945, -</u>	
Fee per additional invention number of additional inv		
Or,x		
The applicant is informed that, according to Rule 40.2(c), the pa i.e., a reasoned statement to the effect that the international appor that the amount of the required additional fee is excessive.	lication complies with the requirement of unity of invention	
3. X Claim(s) Nos. See attached sheet (3nd Article 17(2)(b) because of defects under Article 17(2)(a) a	have been found to be unsearchable under and therefore have not been included with any invention.	
Name and mailing address of the International Searching Authority European Patent Office, P.B. 5818 Patentlaan 2	Authorized officer	
NL-2280 HV Rijswijk	Véronique Baillou	
orm PCT/ISA/206 (July 1992)	ENSCHREIBEN	
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This International Searching Authority found multiple (groups of) inventions in this international application, as follows:

-1. Claims: 1-43

Immunomodulatory polymers, in particular bacterial polysaccharides, and preparations thereof

2. Claims: 44-61

Computer-assisted methods and systems for in silico analysing, predicting, simulating and selecting immunomodulatory polymer candidates

The problem to be solved is how to provide immunomodulatory polymers. The solution in the case of invention 1 is to isolate and purify polymers, in particular natural polysaccharides of bacterial origin. In the case of invention 2 the approach is theoretical: the solution to the above problem consists of the provision of the primary structure of a polymer, the simulation of its 3D structure and a prediction in silico about its potential immunomodulatory properties. Thus, the special technical features of the two solutions are not the same, and the present application is considered to lack unity (Rule 13.1 and 13.2 PCT) The search of the second invention would imply a significant additional effort.

FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 206

Continuation of Box 3.

Although claims 40-43 are directed to a method of treatment of the human/animal body, the search has been carried out and based on the alleged effects of the compound/composition.

Further defect(s) under Article 17(2)(a):

Continuation of Box 3.

Claims Nos.: 22-39 (partially)

Present claims 22-39 relate to a compound defined by reference to the following parameter(s):

3D conformation in solution and geometric-functional features of the same. The use of these parameters in the present context is considered to lead to a lack of clarity within the meaning of Article 6 PCT. It is impossible to compare the parameters the applicant has chosen to employ with what is set out in the prior art. The lack of clarity is such as to render a meaningful complete search impossible. Consequently, the search has been restricted to the polymers disclosed in claims 1-21 in combination with the description, in particular bacterial polysaccharides.

The applicant's attention is drawn to the fact that claims, or parts of claims, relating to inventions in respect of which no international search report has been established need not be the subject of an international preliminary examination (Rule 66.1(e) PCT). The applicant is advised that the EPO policy when acting as an International Preliminary Examining Authority is normally not to carry out a preliminary examination on matter which has not been searched. This is the case irrespective of whether or not the claims are amended following receipt of the search report or during any Chapter II procedure.

An' to Form PCT/ISA/206 COMMUNICA ... JN RELATING TO THE RESULTS OF THE PARTIAL INTERNATIONAL SEARCH

.ternational Application No PCT/US 01/47251

- 1. The present communication is an Annex to the invitation to pay additional fees (Form PCT/ISA/206). It shows the results of the international search established on the parts of the international application which relate to the invention first mentioned in claims Nos.:
- see 'Invitation to pay additional fees'
 2. This communication is not the international search report which will be established according to Article 18 and Rule 43.
- 3.If the applicant does not pay any additional search fees, the information appearing in this communication will be considered as the result of the international search and will be included as such in the international search report.
- 4.If the applicant pays additional fees, the international search report will contain both the information appearing in this communication and the results of the international search on other parts of the international application for which such fees will have been paid.

Category °	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	US 5 700 787 A (TZIANABOS ARTHUR O ET AL) 23 December 1997 (1997-12-23) abstract column 2, line 56 - line 65 column 3, line 5 - line 22 column 4, line 1 - line 11 column 15, line 1 -column 16, line 4 chemical structures tables	1-17,19, 22-40
X	TZIANABOS A O ET AL: "T cells activated by zwitterionic molecules prevent abscesses induced by pathogenic bacteria." THE JOURNAL OF BIOLOGICAL CHEMISTRY. UNITED STATES 10 MAR 2000, vol. 275, no. 10, 10 March 2000 (2000-03-10), pages 6733-6740, XP002206685 ISSN: 0021-9258 abstract page 6733, right-hand column, line 27 - line 47 figure 1 page 6738, line 49 -page 6739, line 8	1-17,19, 22-40

Special categories of cited documents:

- "A" document defining the general state of the art which is not considered to be of particular relevance
- "E" earlier document but published on or after the international filing date
- "L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified)
- "O" document referring to an oral disclosure, use, exhibition or other means
- P" document published prior to the international filing date but later than the priority date claimed
- "T" later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention
- "X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone
- "Y" document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art.
- "&" document member of the same patent family

Anr 'o Form PCT/ISA/206 COMMUNICA. ON RELATING TO THE RESULTS OF THE PARTIAL INTERNATIONAL SEARCH

ernational Application No PCT/US 01/47251

<u> </u>	ntion) DOCUMENTS CONSIDERED TO BE RELEVANT	16
Category °	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	KALKA-MOLL W M ET AL: "Effect of molecular size on the ability of zwitterionic polysaccharides to stimulate cellular immunity." JOURNAL OF IMMUNOLOGY (BALTIMORE, MD.: 1950) UNITED STATES 15 JAN 2000, vol. 164, no. 2, 15 January 2000 (2000-01-15), pages 719-724, XP002206686 ISSN: 0022-1767 abstract; figures 1,5 page 723, last paragraph	1-17,19, 22-43
X .	BRUBAKER J O ET AL: "Mitogenic activity of purified capsular polysaccharide A from Bacteroides fragilis: differential stimulatory effect on mouse and rat lymphocytes in vitro." JOURNAL OF IMMUNOLOGY (BALTIMORE, MD.: 1950) UNITED STATES 15 FEB 1999, vol. 162, no. 4, 15 February 1999 (1999-02-15), pages 2235-2242, XP002206687 ISSN: 0022-1767 page 2235, right-hand column, line 1 - line 7 page 2235, right-hand column, last line -page 2236, left-hand column, line 3 figures 8,9	1-17,19, 22-43
P,A	WANG Y ET AL: "Structural basis of the abscess-modulating polysaccharide A2 from Bacteroides fragilis." PROCEEDINGS OF THE NATIONAL ACADEMY OF SCIENCES OF THE UNITED STATES OF AMERICA. UNITED STATES 5 DEC 2000, vol. 97, no. 25, 5 December 2000 (2000-12-05), pages 13478-13483, XP002206688 ISSN: 0027-8424 the whole document	1-43

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Patent Family Annex

Information on patent family members

...ternational Application No PCT/US 01/47251

Patent document cited in search report	Publication date	Patent family member(s)	Publication date
US 5700787	A 23-12-1997	US 5679654 A AU 1321799 A AU 702325 B2 AU 3543995 A CA 2197563 A1 EP 0778780 A1 JP 10507746 T WO 9607427 A1	21-10-1997 18-03-1999 18-02-1999 27-03-1996 14-03-1996 18-06-1997 28-07-1998 14-03-1996